The claimed invention is:

## 1. A compound of formula (A):

$$\mathbb{R}^{1}$$
 $\mathbb{R}^{4}$ 
 $\mathbb{R}^{3}$ 
 $\mathbb{R}^{3}$ 
 $\mathbb{R}^{3}$ 
 $\mathbb{R}^{4}$ 
 $\mathbb{R}^{4}$ 
 $\mathbb{R}^{4}$ 
 $\mathbb{R}^{4}$ 
 $\mathbb{R}^{4}$ 

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or a pharmaceutically acceptable salt, prodrug, tautomer, hydrate or solvate thereof, wherein:

X is O or S;

R<sup>1</sup> is a saturated, unsaturated, or aromatic C<sub>3</sub>-C<sub>20</sub> mono-, bi- or polycyclic 10 ring optionally containing at least one heteroatom selected from the group consisting of N, O and S, wherein R<sup>1</sup> can optionally be further independently substituted with at least one moiety independently selected from the group consisting of: carbonyl, halo, halo(C<sub>1</sub>-C<sub>6</sub>)alkyl, perhalo(C<sub>1</sub>-C<sub>6</sub>)alkyl, perhalo(C<sub>1</sub>-C<sub>6</sub>)alkoxy, (C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>2</sub>-C<sub>6</sub>)alkenyl, (C<sub>2</sub>-C<sub>6</sub>)alkynyl, hydroxy, oxo, mercapto, (C<sub>1</sub>-15 C<sub>6</sub>)alkylthio, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, (C<sub>5</sub>-C<sub>10</sub>)aryl or (C<sub>5</sub>-C<sub>10</sub>)heteroaryl, (C<sub>5</sub>-C<sub>10</sub>)aryloxy or  $(C_5-C_{10})$ heteroaryloxy,  $(C_5-C_{10})$ ar $(C_1-C_6)$ alkyl or  $(C_5-C_{10})$ heteroar $(C_1-C_6)$ alkyl,  $(C_5-C_{10})$ ar $(C_1-C_6)$ alkoxy or  $(C_5-C_{10})$ heteroar $(C_1-C_6)$ alkoxy, HO-(C=O)-, ester, amido, ether, amino, amino(C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>1</sub>-C<sub>6</sub>)alkylamino(C<sub>1</sub>-C<sub>6</sub>)alkyl,  $di(C_1-C_6)alkylamino(C_1-C_6)alkyl, (C_5-C_{10})heterocyclyl(C_1-C_6)alkyl, (C_1-C_6)alkyl-and$ 20  $di(C_1-C_6)alkylamino$ , cyano, nitro, carbamoyl,  $(C_1-C_6)alkylcarbonyl$ , (C<sub>1</sub>-C<sub>6</sub>)alkoxycarbonyl, (C<sub>1</sub>-C<sub>6</sub>)alkylaminocarbonyl, di(C<sub>1</sub>-C<sub>6</sub>)alkylaminocarbonyl, (C<sub>5</sub>-C<sub>10</sub>)arylcarbonyl, (C<sub>5</sub>-C<sub>10</sub>)aryloxycarbonyl,  $(C_1-C_6)$ alkylsulfonyl, and  $(C_5-C_{10})$ arylsulfonyl;

each R<sup>3</sup> is independently selected from the group consisting of: hydrogen, halo, halo( $C_1$ - $C_6$ )alkyl, ( $C_1$ - $C_6$ )alkyl, ( $C_2$ - $C_6$ )alkenyl, ( $C_2$ - $C_6$ )alkynyl, perhalo(C<sub>1</sub>-C<sub>6</sub>)alkyl, phenyl, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl, hydroxy, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, perhalo(C<sub>1</sub>-C<sub>6</sub>)alkoxy, phenoxy, 5 (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-O-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-O-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-O-,  $(C_1-C_6)$ alkyl-S-,  $(C_1-C_6)$ alkyl-SO<sub>2</sub>-,  $(C_1-C_6)$ alkyl-NH-SO<sub>2</sub>-, O<sub>2</sub>N-, NC-, amino,  $Ph(CH_2)_{1-6}HN-$ ,  $(C_1-C_6)alkyl HN-$ ,  $(C_1-C_6)alkylamino$ ,  $[(C_1-C_6)alkyl]_2$ -amino,  $(C_1-C_6)$ alkyl-SO<sub>2</sub>-NH-, amino(C=O)-, aminoO<sub>2</sub>S-,  $(C_1-C_6)$ alkyl-(C=O)-NH-,  $(C_1-C_6)alkyl-(C=O)-[(((C_1-C_6)alkyl)-N]-, phenyl-(C=O)-NH-,$ phenyl- $(C=O)-[((C_1-C_6)alkyl)-N]-, (C_1-C_6)alkyl-(C=O)-, phenyl-(C=O)-,$ 10  $(C_5-C_{10})$ heteroaryl-(C=O)-,  $(C_5-C_{10})$ heterocyclic-(C=O)-,  $(C_3-C_{10})$ cycloalkyl-(C=O)-, HO-(C=O)-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-O-(C=O)-, H<sub>2</sub>N(C=O)-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-NH-(C=O)-,  $[(C_1-C_6)alkyl]_2-N-(C=O)-$ , phenyl-NH-(C=O)-, phenyl- $[((C_1-C_6)alkyl)-N]-(C=O)-$ , (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-NH-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-NH-(C=O)-, 15  $(C_3-C_{10})$ cycloalkyl-NH-(C=O)- and  $(C_1-C_6)$ alkyl-(C=O)-O-;

where alkyl, alkenyl, alkynyl, phenyl, heteroaryl, heterocyclic, cycloalkyl, alkoxy, phenoxy, amino of  $R^3$  is optionally substituted by at least one substituent independently selected from (C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, halo(C<sub>1</sub>-C<sub>6</sub>)alkyl, halo, H<sub>2</sub>N-, Ph(CH<sub>2</sub>)<sub>1-6</sub>HN-, and (C<sub>1</sub>-C<sub>6</sub>)alkylHN-;

s is an integer from one to five; and

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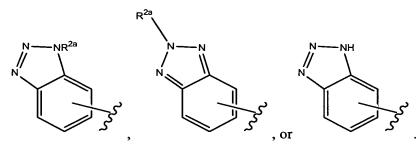
R<sup>4</sup> is selected from the group consisting of: hydrogen, halo, halo(C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>2</sub>-C<sub>6</sub>)alkenyl, (C<sub>2</sub>-C<sub>6</sub>)alkynyl, perhalo(C<sub>1</sub>-C<sub>6</sub>)alkyl, phenyl, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl, hydroxy, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, perhalo(C<sub>1</sub>-C<sub>6</sub>)alkoxy, phenoxy, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-O-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-O-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-O-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-S-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sub>2</sub>-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-NH-SO<sub>2</sub>-, O<sub>2</sub>N-, NC-, amino, Ph(CH<sub>2</sub>)<sub>1-6</sub>NH-, alkylNH-, (C<sub>1</sub>-C<sub>6</sub>)alkylamino, [(C<sub>1</sub>-C<sub>6</sub>)alkyl]<sub>2</sub>-amino, (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sub>2</sub>-NH-, amino(C=O)-, aminoSO<sub>2</sub>-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-NH-,

(C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-((C<sub>1</sub>-C<sub>6</sub>)alkyl)-N]-, phenyl-(C=O)-NH-, phenyl-(C=O)-((C<sub>1</sub>-C<sub>6</sub>)alkyl)-N]-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-, phenyl-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl-(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-(C=O)-, cycloalkyl-(C=O)-, HO-(C=O)-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-O-(C=O)-, H<sub>2</sub>N(C=O)-, (C<sub>1</sub>-C<sub>6</sub>)alkyl-NH-(C=O)-, ((C<sub>1</sub>-C<sub>6</sub>)alkyl)<sub>2</sub>-N-(C=O)-, phenyl-NH-(C=O)-, phenyl-((C<sub>1</sub>-C<sub>6</sub>)alkyl)-N]-(C=O)-, (C<sub>2</sub>-C<sub>10</sub>)heterocyclic-NH-(C=O)-

 $(C_5-C_{10})$ heteroaryl-NH-(C=O)-,  $(C_5-C_{10})$ heterocyclic-NH-(C=O)-,  $(C_3-C_{10})$ cycloalkyl-NH-(C=O)- and  $(C_1-C_6)$ alkyl-(C=O)-O-;

where alkyl, alkenyl, alkynyl, phenyl, heteroaryl, heterocyclic, cycloalkyl, alkoxy, phenoxy, and amino of R<sup>4</sup> is optionally substituted by at least one substituent independently selected from the group consisting of (C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, halo(C<sub>1</sub>-C<sub>6</sub>)alkyl, halo, H<sub>2</sub>N-, Ph(CH<sub>2</sub>)<sub>1-6</sub>-NH-, and (C<sub>1</sub>-C<sub>6</sub>)alkylNH-.

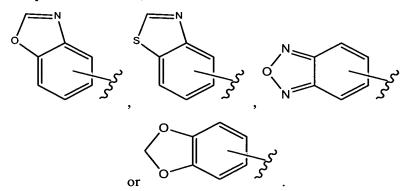
## 2. A compound of claim 1, wherein $R^1$ is



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## 3. A compound of claim 1, wherein $R^1$ is



4. A compound of claim 1, wherein R<sup>1</sup> is

5. A compound of claim 1, wherein  $R^1$  is

6. A compound of claim 1, wherein R<sup>1</sup> is

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7. A compound of claim 1, wherein  $R^1$  is

$$\mathbb{R}^{2a}$$

N

 $\mathbb{R}^{2a}$ 
 $\mathbb{R}^{2a}$ 
 $\mathbb{R}^{2a}$ 
 $\mathbb{R}^{2a}$ 
 $\mathbb{R}^{2a}$ 
 $\mathbb{R}^{2a}$ 
 $\mathbb{R}^{2a}$ 

8. A compound of claim 1, wherein  $R^1$  is

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- 9. A compound of claim 1, wherein X is O; s is one to two;  $R^3$  is hydrogen or  $(C_1-C_6)$ alkyl; and  $R^4$  is H,  $(C_1-C_6)$ alkyl, or  $(C_3-C_{10})$ cycloalkyl.
- 10. A compound of claim 1, wherein X is S; s is one to two;  $R^3$  is hydrogen or  $(C_1-C_6)$ alkyl; and  $R^4$  is H,  $(C_1-C_6)$ alkyl, or  $(C_3-C_{10})$ cycloalkyl.
- 11. A pharmaceutical composition comprising a compound of claim 1 and a15 pharmaceutically acceptable carrier.
  - 12. A method of preventing or treating a TGF-related disease state in an animal or human comprising the step of administering a therapeutically effective amount of

a compound of claim 1 to the animal or human suffering from the TGF-related disease state.

13. A method of claim 12, wherein said TGF-related disease state is selected
 5 from the group consisting of cancer, glomerulonephritis, diabetic nephropathy,
 hepatic fibrosis, pulmonary fibrosis, intimal hyperplasia and restenosis, scleroderma,
 and dermal scarring.

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